

AMENDMENT AND RESPONSE

Serial Number: 09/150,813

Filing Date: September 11, 1998

Title: COMPOUNDS AND METHODS TO INHIBIT OR AUGMENT AN INFLAMMATORY RESPONSE

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neutrophil activating protein-2 (NAP-2) [NAP-2].

B3  
20. (Amended) A method of preventing or inhibiting an indication associated with hematopoietic cell recruitment, comprising: administering to a mammal at risk of, or afflicted with, the indication an effective amount of a [chemokine peptide 3] peptide of a chemokine, a variant thereof, a derivative thereof, or a combination thereof, wherein the peptide comprises no more than 30 amino acid residues, wherein at least three contiguous residues of the peptide correspond to residues in the carboxyl-terminal half of the mature form of the chemokine, wherein the three contiguous residues correspond to residues Trp-Val-Gln or Lys-Gln-Lys in human MCP-1, and wherein the peptide inhibits the response induced by the corresponding native chemokine[, a chemokine peptide 2, a variant thereof, a derivative thereof, a compound of formula (IV), a compound of formula (V), a compound of formula (VI), or a combination thereof].

B4  
22. (Amended) A method to modulate the chemokine-induced activity of hematopoietic cells at a preselected physiological site, comprising: administering to a mammal a dosage form comprising an effective amount of a [chemokine peptide 3] peptide of a chemokine, a variant thereof, a derivative thereof, or a combination thereof, wherein the peptide comprises no more than 30 amino acid residues, wherein at least three contiguous residues of the peptide correspond to residues in the carboxyl-terminal half of the mature form of the chemokine, wherein at least three contiguous residues of the peptide correspond to residues in the carboxyl-terminal half of the mature form of the chemokine, wherein the three contiguous residues correspond to residues Trp-Val-Gln or Lys-Gln-Lys in human MCP-1, and wherein the peptide inhibits the response induced by the corresponding native chemokine, [a chemokine peptide 2, a variant thereof, a derivative thereof, a compound of formula (IV), a compound of formula (V), a compound of formula (VI), or a combination thereof,] wherein the dosage form is linked to a site targeting moiety.

B5  
34. (Amended) A method to [increase or enhance] ~~alter~~ hematopoietic cell-associated activity at a tumor site, comprising: administering an effective amount of a [chemokine peptide

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3] peptide of a chemokine, a variant thereof, a derivative thereof, or a combination thereof, wherein the peptide comprises no more than 30 amino acid residues, and wherein at least three contiguous residues of the peptide correspond to residues in the carboxyl-terminal half of the mature form of the chemokine, wherein the three contiguous residues correspond to residues Trp-Val-Gln or Lys-Gln-Lys in human MCP-1], a chemokine peptide 2, a variant thereof, a derivative thereof, a compound of formula (IV), a compound of formula (V), a compound of formula (VI), or a combination thereof].

Please add the following new claims:

52. (New) The method of claim 17, 20, 22 or 34, wherein the peptide of a chemokine is a peptide of a CC chemokine.

53. (New) The method of claim 52, wherein the CC chemokine is monocyte chemotactic protein-1 (MCP-1), regulated on activation, normal T expressed and secreted protein (RANTES), monocyte chemotactic protein-2 (MCP-2), monocyte chemotactic protein-3 (MCP-3), monocyte chemotactic protein-4 (MCP-4), eotaxin, macrophage inflammatory protein-1 $\alpha$  (MIP1 $\alpha$ ), MIP1 $\beta$ , liver and activation regulated chemokine (LARC), I309, hemofiltrate CC-chemokine -1 (HCC-1), thymus and activation regulated chemokine (TARC) or chemokine beta 8 (Ck $\beta$ 8).

54. (New) The method of claim 17, 20, 22 or 34, wherein the peptide of a chemokine is a peptide of a CXC chemokine.

55. (New) The method of claim 20, 22 or 34, wherein the CXC chemokine is interleukin 8 (IL-8), interferon inducible protein 10 (IP-10), platelet factor-4 (PF-4), stromal cell-derived factor-1 (SDF-1 $\alpha$ ), neutrophil activating protein-2 (NAP-2), growth regulated oncogene alpha (GRO $\alpha$ ), GRO $\beta$ , GRO $\gamma$  or epithelial neutrophil activating peptide-78 (ENA78).

56. (New) The method of claim 17, wherein the CXC chemokine is interferon inducible protein 10 (IP-10), platelet factor-4 (PF-4), stromal cell-derived factor-1 (SDF-1 $\alpha$ ), growth regulated oncogene alpha (GRO $\alpha$ ), GRO $\beta$ , GRO $\gamma$  or epithelial neutrophil activating peptide-78

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*See  
C1  
cont*  
(ENA 78).

57. (New) The method of claim 56, wherein the variant peptide is Glu-Ile-Cys-Leu-Asp-Pro-Lys-Gln-Lys-Trp-Ile-Gln (SEQ ID NO:14).

58. (New) The method of claim 17, 20, 22 or 34, wherein the peptide of a chemokine comprises SEQ ID NO:1, SEQ ID NO:7, SEQ ID NO:38, SEQ ID NO:40, SEQ ID NO:41, SEQ ID NO:42, SEQ ID NO:43, SEQ ID NO:44, SEQ ID NO:65, SEQ ID NO:66, SEQ ID NO:67, SEQ ID NO:68, SEQ ID NO:72, SEQ ID NO:73, or SEQ ID NO:74.

*B6*  
59. (New) The method of claim 17, 20, 22, or 34, wherein the peptide of a chemokine is a cyclic reverse D sequence (CRD) derivative or a variant thereof.

60. (New) The method of claim 59, wherein the CRD derivative is CRD-Cys-Leu-Asp-Pro-Lys-Gln-Lys-Trp-Ile-Gln-Cys.

61. (New) The method of claim 17, 20, 22, or 34, wherein the variant peptide is a variant peptide of peptide 3(3-12)[MCP-1].

62. (New) The method of claim 17, wherein the indication is atherosclerosis, multiple sclerosis, hypertension, asthma, allergy, psoriasis, rheumatoid arthritis, osteoporosis, stroke, acute ischemia, or organ transplant rejection.

**Remarks**

Reconsideration and withdrawal of the rejections of the claims, in view of the amendments and remarks presented herein, is respectfully requested. Claims 17, 20, 22, and 34 are amended, claims 1-16, 18-19, 23, 29-30, 36-39, 46-47, and 51 are canceled, and claims 52-62 are added. Claims 17, 20-22, 24-28, 31-35, 40-45, 48-50, and 52-62 are pending. The amendments to the claims are intended to clarify Applicant's invention and are not intended to limit the equivalents to which any claim is entitled.